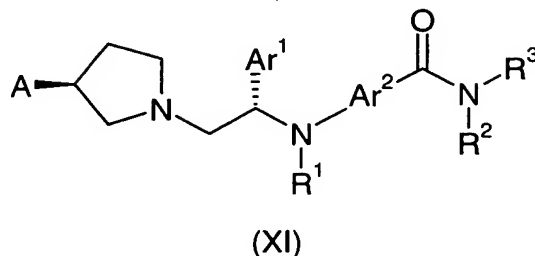


Claims

1. A single-step or multi-step process for the preparation of a compound of formula (XI):



or a stereoisomer thereof, wherein;

10 A is hydrogen, hydroxy, C₁-C₆ (preferably C₁-C₄) alkyl, C₁-C₆ (preferably C₁-C₄) fluoroalkyl (particularly -CF₃), C₁-C₆ (preferably C₁-C₄) alkoxy, or OY wherein Y is a hydroxy protecting group or A, taken together with its geminal hydrogen, is an oxo group;

15 Ar¹ is phenyl optionally substituted by one or more (preferably one to two) substituents selected from fluoro, C₁-C₄ alkyl, C₁-C₄ alkoxy, C₁-C₄ alkoxy-C₁-C₄ alkoxy, trifluoromethyl, carboxy-C₁-C₄ alkoxy and C₁-C₄ alkoxycarbonyl-C₁-C₄ alkoxy;

20 Ar² is phenyl, naphthyl, pyridyl, thienyl, furyl, pyrrolyl or pyrimidyl, each being optionally substituted by one or more (preferably one to two) substituents selected from fluoro, C₁-C₄ alkyl, C₁-C₄ alkoxy, di(C₁-C₄)alkylamino and C₁-C₄ fluoroalkyl;

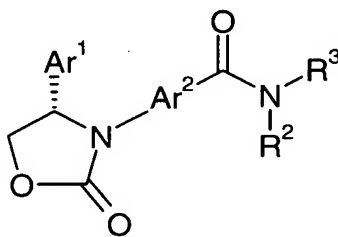
25 R¹ is C₁-C₆ alkyl or benzyl wherein the phenyl moiety of said benzyl is optionally substituted with C₁-C₆ alkoxy or OY wherein Y is a hydroxy protecting group; and

R^2 and R^3 are independently selected from hydrogen, C_1 - C_7 alkyl optionally substituted by one or more (preferably one to five) hydroxy or halo groups, C_3 - C_6 cycloalkyl, C_2 - C_6 alkenyl, C_2 - C_6 alkynyl, C_1 - C_7 (preferably C_1 - C_5) alkoxy, phenyl optionally substituted by fluoro (preferably substituted by one or two fluoro groups), phenyl- C_1 - C_7 (preferably C_1 - C_5) alkyl wherein the phenyl group is optionally substituted by fluoro, and $-(CH_2)_nX-R^4$ wherein n is one or two, X is O or S and R^4 is C_1 - C_3 alkyl, or, when Ar^2 is phenyl, $-Ar^2-C(=O)-N(R^2)-$ is a phthalimide group and R^3 is C_1 - C_7 alkyl; or

R^2 and R^3 , together with the nitrogen atom to which they are attached, form a pyrrolidine, piperidine or morpholine ring, optionally substituted by C_1 - C_3 alkyl or fluoro;

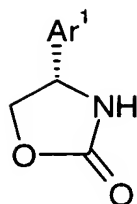
comprising a step in which the $N-Ar^2$ bond is constructed by a copper-mediated aryl amination.

2. A process as claimed in claim 1 wherein a compound of formula (IV):



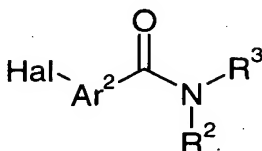
(IV)

or the enantiomer thereof, wherein Ar^1 , Ar^2 , R^2 and R^3 are as defined in claim 1, is prepared by treating a compound of formula (II):



(II)

or the enantiomer thereof, wherein Ar¹ is as defined in claim 1, with a
5 compound of formula (III):



(III)

10 wherein Ar², R² and R³ are as defined in claim 1 and wherein one unsubstituted position on the Ar² moiety is substituted with a halogen group Hal, preferably Cl, Br or I, most preferably Br, in the presence of a cuprous salt, an amino ligand and a base.

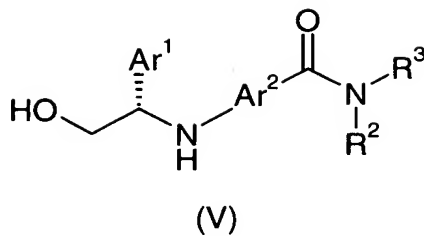
15 3. A process as claimed in claim 2 wherein the cuprous salt is CuI, CuBr or CuCl.

4. A process as claimed in claim 2 wherein the amino ligand is 1,2-diaminocyclohexane.

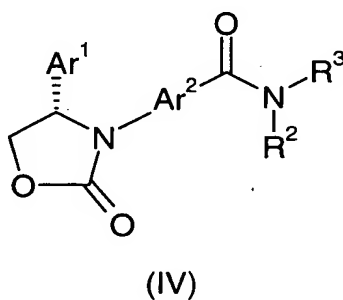
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5. A process as claimed in claim 2 wherein the base is sodium carbonate, potassium carbonate or cesium carbonate.

6. A process as claimed in claim 1 wherein a compound of formula
25 (V):

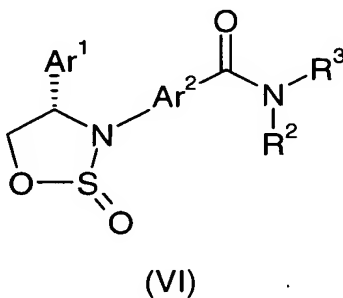


or the enantiomer thereof, wherein Ar¹, Ar², R² and R³ are as defined in
 5 claim 1, is prepared by treating a compound of formula (IV):

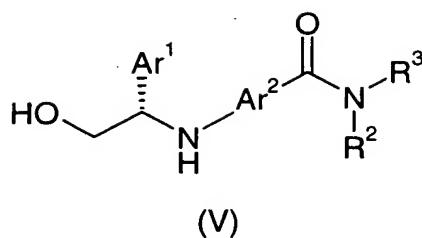


10 or the enantiomer thereof, wherein Ar¹, Ar², R² and R³ are as defined in
 claim 1, with a base in the presence of water.

7. A process as claimed in claim 1 wherein a compound of formula
 formula (VI):

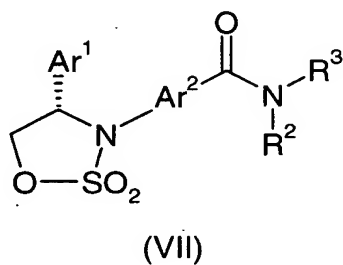


wherein Ar¹, Ar², R² and R³ are as defined in claim 1, or the enantiomer
 20 thereof, is prepared by treating a compound of formula (V):

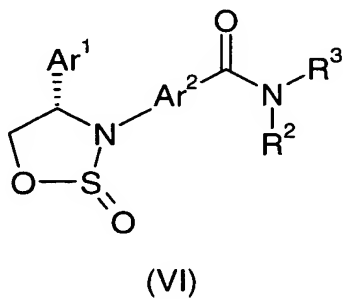


5 or the enantiomer thereof, wherein Ar¹, Ar², R² and R³ are as defined in claim 1, with a thionyl halide.

8. A process as claimed in claim 1 wherein a compound of formula
10 (VII):



15 wherein Ar¹, Ar², R² and R³ are as defined in claim 1, or the enantiomer thereof, is prepared by oxidising a compound of formula (VI):



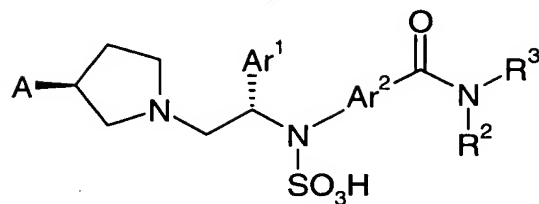
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wherein Ar¹, Ar², R² and R³ are as defined in claim 1, or the enantiomer

thereof.

9. A process as claimed in claim 1 wherein a compound of formula (IX):

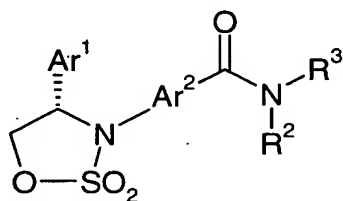
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(IX)

wherein A, Ar¹, Ar², R² and R³ are as defined in claim 1, or a zwitterion thereof, or a stereoisomer of either, is prepared by treating a compound of formula (VII):

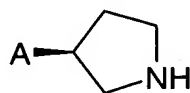
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(VII)

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wherein Ar¹, Ar², R² and R³ are as defined in claim 1, or the enantiomer thereof, with a compound of formula (VIII):

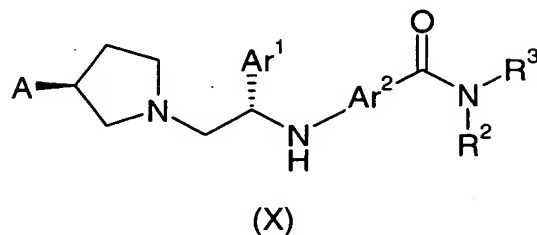


(VIII)

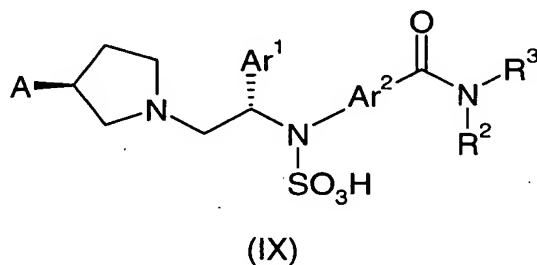
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wherein A is as defined in claim 1, or the enantiomer thereof.

10. A process as claimed in claim 1 wherein a compound of formula (X):



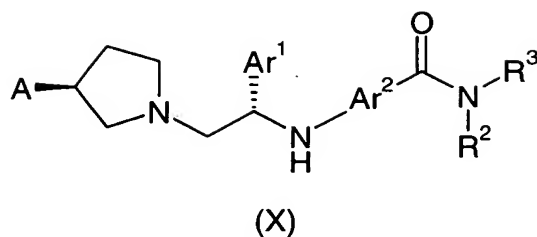
wherein A, Ar¹, Ar², R² and R³ are as defined in claim 1, or a stereoisomer thereof is prepared by hydrolytically cleaving the -SO₃H group in a compound of formula (IX):



15 wherein A, Ar¹, Ar², R² and R³ are as defined in claim 1, or a zwitterion thereof, or a stereoisomer of either.

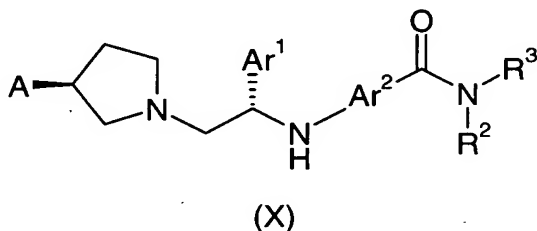
11. A process as claimed in claim 1 wherein a compound of the formula (XI), as defined in claim 1, or a stereoisomer thereof, is prepared by the reductive alkylation of a compound of formula (X):

20



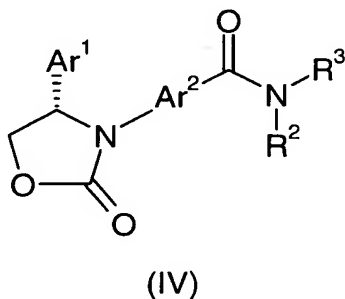
wherein A, Ar¹, Ar², R² and R³ are as defined above, or a stereoisomer thereof.

12. A process for the preparation of a compound of formula (XI), as defined in claim 1, or a stereoisomer thereof, comprising the reductive amination of a compound of formula (X):



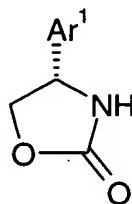
or a stereoisomer thereof, wherein A, Ar¹, Ar², R² and R³ are as defined in claim 1.

13. A process for the preparation of a compound of formula (IV):



or the enantiomer thereof, wherein Ar¹, Ar², R² and R³ are as defined in

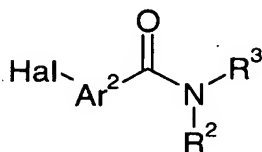
claim 1, comprising treating a compound of formula (II):



(II)

5

or the enantiomer thereof, wherein Ar¹ is as defined in claim 1, with a compound of formula (III):



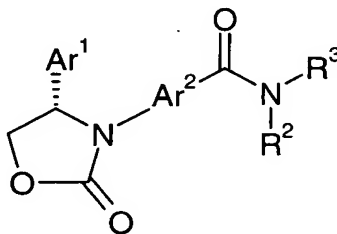
(III)

10

wherein Ar², R² and R³ are as defined in claim 1 and wherein one unsubstituted position on the Ar² moiety is substituted with a halogen group Hal, preferably Cl, Br or I, most preferably Br, in the presence of a cuprous salt, an amino ligand and a base.

15

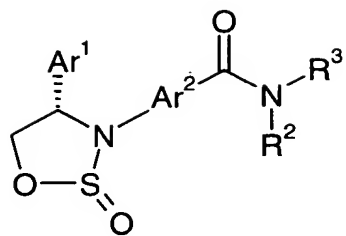
14. A compound of formula:



(IV)

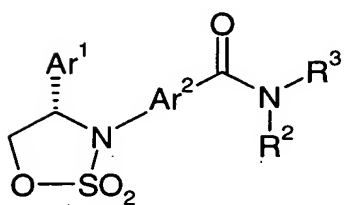
20

or



(VI)

or



(VII)

wherein Ar¹, Ar², R² and R³ are as defined in claim 1.